Human TCR Vα-1.5 (Vα-8.2) coding sequence

AACCAGAGCC CAGTCGGTGA CCCAGCTTGA CAGCCACGTC TCTGTCTCTG GAAGTACACA TCAGCGGCCA CCCTGGTTAA AGGCATCAAC GGTTTTGAGG CATATGAGCG ACGCGGCTGA GTACTTCTGT GTTGTGAGTC CTTTTTCAGG TGCTAGACAT GAGGTCTATG GACTTCAAGA GCAACAGTGC TGTGGCCTGG ATGCTCCTGC TGCTCGTCCC AGTGCTCGAG GTGATTTTTA CTCTGGGAGG AGGAGGTGCT GACGGACTCA CCTTTGGCAA AGGGACTCAT CTAATCATCC AGCCCTATAT CCAGAACCCT GACCCTGCCG TGTACCAGCT GAGAGACTCT CTGAATTTAA GAAGAGTGAA ACCTCCTTCC ACCTGACGAA ACCCTCAGCC TCCAGAGAC ACCTTCTTCC CCAGCCCAGA AAGTTCCTGT GATGTCAAGC AAGGAACCCC GGTGCTGCTG AGGTGCAACT ACTCATCTTC TTATTCACCA ICAGTGATTG GGTTCCGAAT CCTCCTCCTG AAAGTGGCCG GGTTTAATCT FGGTCGAGAA AAGCTTTGAA ACAGATACGA ACCTAAAACTT TCAAAACCTG TCTCTCTTCT GGTATGTGCA ACACCCCAAC AAAGGACTCC AGCTTCTCCT AAATGTGTCA CAAAGTAAGG ATTCTGATGT GTATATCACA GACAAACTG AAATCCAGTG ACAAGTCTGT CTGCCTATTC ACCGATTTTG ATTCTCAAAC AGCAACAAAT CTGACTTTGC ATGTGCAAAC GCCTTCAACA ACAGCATTAT GCTCATGACG CTGCGGCTGT GGTCCAGCTG A

Figure 1:

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MLLLLVPVLEVIFTLGGTRAQSVTQLDSHVSVSEGT Human TCR V α -1.5 (V α -8.2) protein sequence

PVLLRCNYSSSYSPSLFWYVQHPNKGLQLLLKYT

CDR2

FR3

SAATLVKGINGFEAEFKKSETSFHLTKPSAHMSDA

CDR3

AEYFCVVSPFSGGGADGLT

IFFPSPESSCDVKLVEKSFETDTNLNFQNLSVIGFRIL FGKGTH LIIQPYIQNP DPAVYQLRDSKSSDKSVCLF TDFDSQTNVS QSKDSDVYIT DKTVLDMRSM **JEKSNSAVAWSNKSDFACAN AFNNSIPED** LL KVAGFNLLMT LRLWSS constant

Human TCR V β -2.1 (V β -20.1) coding sequence

GGAGACCCAG TACTTCGGGC CAGGCACGCG GCTCCTGGTG CTCGAGGACC CTACCCCGAC CACGTGGAGC TGAGCTGGTG GGTGAATGGG AAGGAGGTGC ACAGTGGGGT CAGCACAGAC CCGCAGCCCC TCAAGGAGCA GCCCGCCCTC ATCGTCAGCG CCGAGGCCTG GGGTAGAGCA GACTGTGGCT TCACCTCCGA ICTCGGAGAA TGACGAGTGG ACCCAGGATA GGGCCAAACC TGTCACCCAG FAGGGAAGGC CACCTTGTAT GCCGTGCTGG TCAGTGCCCT CGTGCTGATG CTGGCAGAAC CCCCGCAACC ACTTCCGCTG TCAAGTCCAG TTCTACGGGC ATGCTGCTGC TTCTGCTGCT TCTGGGGCCA GGCTCCGGGC TTGGTGCTG1 CTCCAAGGCC ACATACGAGC AAGGCGTCGA GAAGGACAAG TTTCTCATCA TGAAAAACGT GTTCCCACCC GAGGTCGCTG TGTTTGAGCC ATCAGAAGCA GAGATCTCCC ACACCCAAAA GGCCACACTG GTGTGCCTGG CCACAGGCTT CGTCAGTTCC CGAAACAGAG TCTCATGCTG ATGGCAACTT CCAATGAGGG GAAGACAGCA GCTTCTACAT CTGCAGTGCT AGAGATGGGG GGGAGGGTT(AATGACTCCA GATACTGCCT GAGCAGCCGC CTGAGGGTCT CGGCCACCTI CGTCTCTCAA CATCCGAGCT GGGTTATCTG TAAGAGTGGA ACCTCTGTGA ACCATGCAAG CCTGACCTTG TCCACTCTGA CAGTGACCAG TGCCCATCCT GTCTTACCAG CAAGGGGTCC TGTCTGCCAC CATCCTCTAT GAGATCTTGC AGATCGAGTG CCGTTCCCTG GACTTTCAGG CCACAACTAT GTTTTGGTAT SCCATGGTCA AGAGAAAGGA TTCCAGAGGC TAG

MLLLLLLGPGSGLGAVVSQHPSWVICKSGTSVKIECR

Human TCR V β -2.1 (V β -20.1) protein sequence

Figure 4

CDR1

FR2

SLDFQATTMFWYRQFPKQSLMLMATSNEGSKATYEQ

GVEKDKFLINHASLTLSTLTVTSAHPEDSSFYICSARD

GGEG CDR3

constant

KATLVCLATGFYPDHVELSWWVNGKEVHSGVSTDPQPL GLSENDEWTQDRAKPVTQIVSAEAWGRADCGFTSESYQ **QGVLSATILYEILLGKATLYAVLVSALVLMAMVKRKDS** KEOPALNDSRYCLSSRLRVSATFWONPRNHFRCQVQFY SETQYFGPGTRLLVLEDLKNVFPPEVAVFEPSEAEISHTQ

Figure 5

Human TCR V α -1.5 (V α -8.2) protein sequence

MILLLVPVLEVIFTLGGTRAQSVTQLDSHVSVSEGT

CDR1

PVLLRCNYSSSYSPSLFWYVQHPNKGLQLLLKYT

FR3

SAATLVKGINGFEAEFKKSETSFHLTKPSAHMSDA CDR2

145

CDR3

Va8.2

AEYFCVVSPFSGGGADGLTFGKGTH LIIQP

constant

TFFPSPESSCDVKLVEKSFETDTNLNFQNLSVIGFRIL YIQNP DPAVYQLRDSKSSDKSVCLF TDFDSQTNVS **JEKSNSAVAWSNKSDFACAN AFNNSIPED OSKDSDVYIT DKTVLDMRSM** LL KVAGFNLLMT LRLWSS

Human TCR V β -2.1 (V β -20.1) protein sequence

figure 6

FR1

MILLILLGPGSGLGAVVSQHPSWVICKSGTSVKTECR

SLDFQATTMFWYRQFPKQSLMLMATSNEGSKATYEQ FR2

GVEKDKFLINHASLTLSTLTVTSAHPEDSSFYICSARD

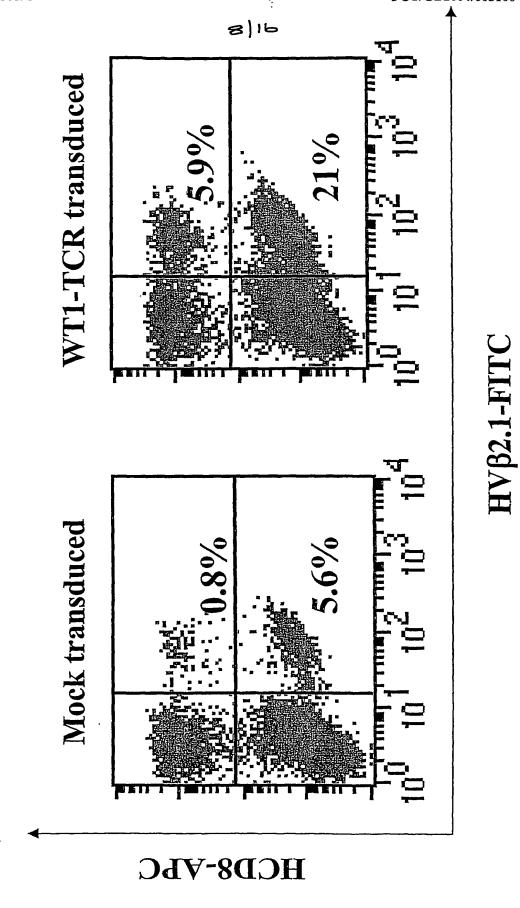
CDR3 GGEGSETOYFGPGTRLLVL

RLRVSATFWQNPRNHFRCQVQFYGLSENDEWTQDRAKP EDLKNVFPPEVAVFEPSEAEISHTOKATLVCLATGFYPDH VELSWWVNGKEVHSGVSTDPQPLKEQPALNDSRYCLSS VTQIVSAEAWGRADCGFTSESYQQGVLSATILYEILLGK ATLYAVLVSALVLMAMVKRKDSRG Constant 2

WO 2005/056595 PCT/GB2004/005100

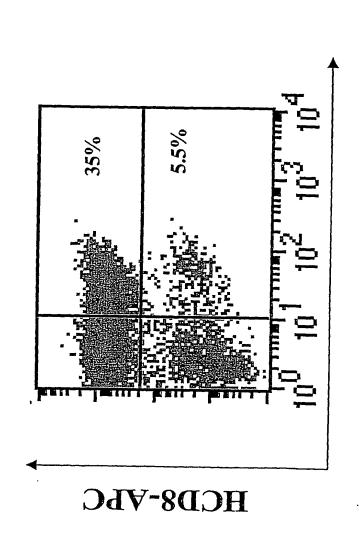
7/16 3'-LTR 3'-LTR PRE TCR-retroviral constructs TCR-pMP71-Gpre 7 kb pMP71-Gpre 7 kb Amp Amp

TCR Expression in Human PBMC after transduction



 ∞

Increase of CD8⁺-Vb2.1⁺ T Cells after antigen stimulation



HVβ2.1-FITC

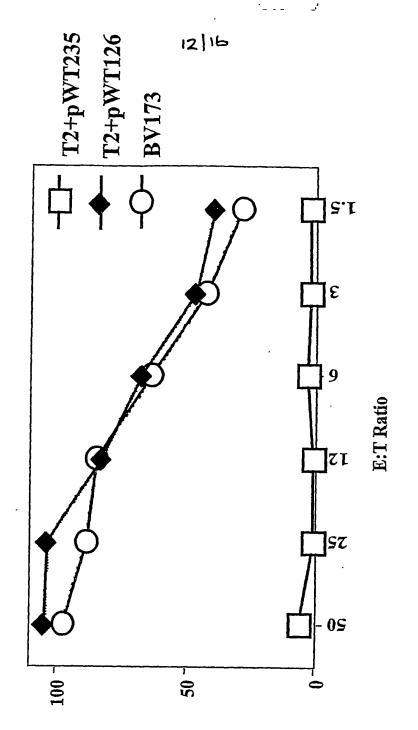
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TCR specific for pWT126 transduced PBMC

igure

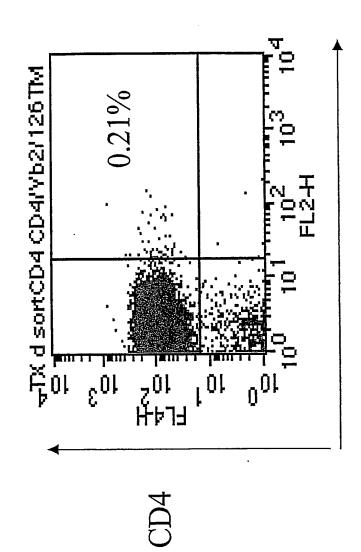
% Shecieic Kitting

TCR transduced CD8+ T cells show pWT126-specific killing activity



% SLECILIC KILLING

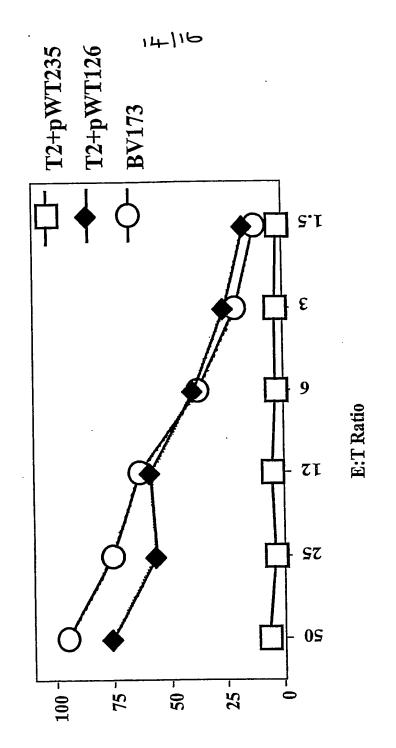
TCR specific for pWT126 transduced PBMC sorted CD4



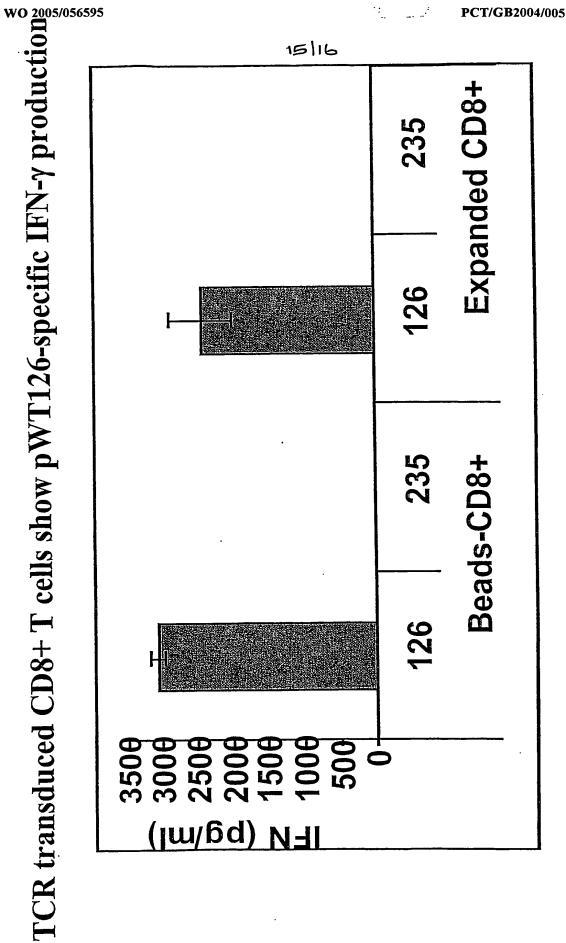
Tetramer

Figure 14

TCR transduced CD4+ T cells show pWT126-specific killing activity



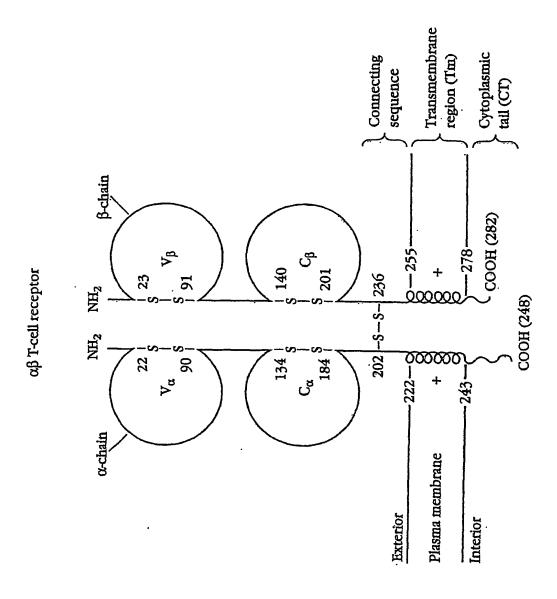
% SLECILIC KILLING



After 20 hrs incubation

WO 2005/056595 PCT/GB2004/005100

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